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INTERNATIONAL SEARCH REPORT

(PCT Article 18 and Rules 43 and 44)

Applicant's or agent's file reference	FOR FURTHER	see Form PCT/ISA/220			
WPP290112	AGIIGII	well as, where applicable, item 5 below. (Earliest) Priority Date (day/month/year)			
International application No.	International filing date (day/month/year)	(Edilest) Priority Data (day/monabyear)			
PCT/GB2005/050042	23/03/2005 23/03/2004				
Applicant					
BABRAHAM INSTITUTE					
This international Search Report has been prepared by this international Searching Authority and is transmitted to the applicant according to Article 18. A copy is being transmitted to the international Bureau.					
This International Search Report consists of a total of05 sheets.					
	a copy of each prior art document cited in t	nis report.			
Basis of the report					
 a. With regard to the language, the language in which it was filed, un 	international search was carried out on the less otherwise indicated under this item.				
this Authority (Ru	ile 23.1(b)).	nslation of the international application furnished to			
b. With regard to any nucle	otide and/or amino acid sequence disclos	ed in the International application, see Box No. I.			
2. Certain claims were fou	und unsearchable (See Box II).				
3. Unity of Invention is lac	eking (see Box III).				
4. With regard to the title,					
I I	ubmitted by the applicant.	<u>—</u>			
the text has been establi	shed by this Authority to read as follows:	BEST AVAILAB			
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5. With regard to the abstract,					
X the text is approved as s	submitted by the applicant.	hority as it appears in Box No. IV. The applicant			
the text has been estable may, within one month f	remed, according to have 36.2(a), by the har- rom the date of mailing of this international	hority as it appears in Box No. IV. The applicant search report, submit comments to this Authority.			
6. With regard to the drawings,					
	published with the abstract is Figure No	01			
as suggested by the applicant.					
as selected by this Authority, because the applicant failed to suggest a figure.					
as selected by this Authority, because this figure better characterizes the invention. b. \(\) none of the figures is to be published with the abstract.					
b none of the figures is to	ne haaltellen waa die dhereme				

Form PCT/ISA/210 (first sheet) (January 2004)

A. CLASSIFICATION OF SUBJECT MATTER IPC 7 G01N33/68

According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols) IPC $\frac{7}{601N}$

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the International search (name of data base and, where practical, search terms used)

EPO-Internal, BIOSIS, WPI Data

Category °	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X,P	WALKER SIMON A ET AL: "Identification of a Ras GTPase-activating protein regulated by receptor-mediated Ca2+ oscillations" EMBO (EUROPEAN MOLECULAR BIOLOGY ORGANIZATION) JOURNAL, vol. 23, no. 8, April 2004 (2004-04), pages 1749-1760, XP002348353 ISSN: 0261-4189 page 1750 - page 1757	1-38

Y Further documents are listed in the continuation of box C.	X Patent family members are listed in whitex.		
Special categories of cited documents: A document defining the general state of the art which is not considered to be of particular relevance E earlier document but published on or after the International filling date L document which may throw doubts on priority claim(e) or which is cited to establish the publication date of another citation or other special reason (as specified) O document referring to an oral disclosure, use, exhibition or other means P document published prior to the international filling date but later than the priority date claimed	"T" later document published siter the International filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention with the principle of the considered to cannot be considered novel or cannot be considered to Involve an Inventive step when the document is taken alone occument of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents; such combined with one or more other such documents, such combined with one or more other such documents, such combination being obvious to a person skilled in the art. "3" document member of the same patent family		
Date of the actual completion of the international search	Date of mailing of the international search report		
10 October 2005	07/11/2005		
Name and mailing address of the ISA	Authorized officer		
European Patent Office, P.B. 5818 Patentiaan 2 NL - 2280 HV Rijswijk Tel. (+31-70) 340-2040, Tx. 31 651 epo nl. Fax: (+31-70) 340-3016	GONCALVES MAUGER, M		

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Citation of document, with indication, where annundate of the missest accounts	
Treatment necessary, where appropriate, or the relevant passages	Relevant to claim No.
BIVONA TREVER G ET AL: "Phospholipase Cgamma activates Ras on the Golgi apparatus by means of RasGRP1." NATURE (LONDON), vol. 424, no. 6949, 7 August 2003 (2003-08-07), pages 694-698, XP002348354 ISSN: 0028-0836 page 696 - page 698	1-38
LOCKYER P J ET AL: "CAPRI REGULATES CA2+-DEPENDENT INACTIVATION OF THE RAS-MAPK PATHWAY" CURRENT SCIENCE, GB, vol. 11, no. 12, 2001, pages 981-986, XP001183507 ISSN: 0960-9822 the whole document	1-38
WALKER S A ET AL: "Control of Ras cycling by Ca<2+>" FEBS LETTERS, ELSEVIER, AMSTERDAM, NL, vol. 546, no. 1, 3 July 2003 (2003-07-03), pages 6-10, XP004433612 ISSN: 0014-5793 page 6 - page 8	1-38
WALKER SIMON A ET AL: "Analyzing the role of the putative inositol 1,3,4,5-tetrakisphosphate receptor GAP1IP4BP in intracellular Ca2+ homeostasis." JOURNAL OF BIOLOGICAL CHEMISTRY, vol. 277, no. 50, 13 December 2002 (2002-12-13), pages 48779-48785, XP002348355 ISSN: 0021-9258 page 48780 - page 48784	1-38
CULLEN P J ET AL: "INTEGRATION OF CALCIUM AND RAS SIGNALLING" NATURE REVIEWS MOLECULAR CELL BIOLOGY, MACMILLAN MAGAZINES, LONDON, GB, vol. 3, no. 5, May 2002 (2002-05), pages 339-348, XP009036294 page 344; figure 3 -/	1-38
	Cgamma activates Ras on the Golgi apparatus by means of RasGRP1." NATURE (LONDON), Vol. 424, no. 6949, 7 August 2003 (2003-08-07), pages 694-698, XP002348354 ISSN: 0028-0836 page 696 - page 698 LOCKYER P J ET AL: "CAPRI REGULATES CA2+-DEPENDENT INACTIVATION OF THE RAS-MAPK PATHWAY" CURRENT BIOLOGY, CURRENT SCIENCE,, GB, Vol. 11, no. 12, 2001, pages 981-986, XP001183507 ISSN: 0960-9822 the whole document WALKER S A ET AL: "Control of Ras cycling by Ca<2+>" FEBS LETTERS, ELSEVIER, AMSTERDAM, NL, Vol. 546, no. 1, 3 July 2003 (2003-07-03), pages 6-10, XP004433612 ISSN: 0014-5793 page 6 - page 8 WALKER SIMON A ET AL: "Analyzing the role of the putative inositol 1,3,4,5-tetrakisphosphate receptor GAPIIPABP in intracellular Ca2+ homeostasis." JOURNAL OF BIOLOGICAL CHEMISTRY, Vol. 277, no. 50, 13 December 2002 (2002-12-13), pages 48779-48785, XP002348355 ISSN: 0021-9258 page 48780 - page 48784 CULLEN P J ET AL: "INTEGRATION OF CALCIUM AND RAS SIGNALLING" NATURE REVIEWS MOLECULAR CELL BIOLOGY, MACMILLAN MAGAZINES, LONDON, GB, Vol. 3, no. 5, May 2002 (2002-05), pages 339-348, XP009036294 page 344; figure 3

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Category °	ation) DOCUMENTS CONSIDERED TO BE RELEVANT Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	LOGAN-SMITH MELANIE J ET AL: "Curcumin, a molecule that inhibits the Ca2+-ATPase of sarcoplasmic reticulum but increases the rate of accumulation of Ca2+" JOURNAL OF BIOLOGICAL CHEMISTRY, vol. 276, no. 50, 14 December 2001 (2001-12-14), pages 46905-46911, XP002348356 ISSN: 0021-9258 abstract	1-38
P,X	WO 2005/003783 A (BABRAHAM INSTITUTE; LOCKYER, PETER) 13 January 2005 (2005-01-13) page 36 - page 44	1-38
A	US 6 514 709 B1 (GRANT STEPHAN K ET AL) 4 February 2003 (2003-02-04) abstract; claims	1-38
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Information on	patent	family	members

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PCT/6	B2005/	050042

Parent document cited in search report		Publication date		Patent family member(s)	Publication date
WO 2005003783		13-01-2005	NONE		
US 6514709	B1	04-02-2003	NONE	و که خدر بی در کارگذاری کارک کارک کارک کارک کارک کارک کارک ک	